

I&W

LAHIVE
&
COCKFIELD
 L L P

OCT. 8 2004
 PATENT & TRADEMARK OFFICE
 COUNSELLORS AT LAW
 28 STATE STREET
 BOSTON, MASSACHUSETTS 02109-1784
 TELEPHONE (617) 227-7400
 FAX (617) 742-4214
 www.lahive.com

JOHN A. LAHIVE, JR. (1928-1997)	MERIDETH C. ARNOLD	PATENT AGENTS
THOMAS V. SMURZYNSKI	DANIELLE L. HERRITT	JONATHAN M. SPARKS, Ph.D.
GIULIO A. DeCONTI, JR.	EUIHOON LEE **	ANDRINA WILLIAMS ZINK
ELIZABETH A. HANLEY	MANEESH GULATI	CRISTIN E. HOWLEY, Ph.D.
AMY BAKER MANDRAGOURAS	CYNTHIA M. SOROS	JILL ANN MELLO, Ph.D.
ANTHONY A. LAURENTANO	PETER W. DINI, Ph.D.	CHRISTOPHER E. DRABIK
KEVIN J. CANNING	MICHAEL J. BASTIAN, Ph.D.	
JANE E. REMILLARD	CHRISTOPHER J. MCKENNA	TECHNICAL SPECIALISTS
DeANN FORAN SMITH	VINCENT P. LOCCISANO	CATHERINE M. BISHOP
DEBRA J. MILASINCIC, Ph.D.	JAMES M. McKENZIE	JACOB G. WEINTRAUB
WILLIAM A. SCOFIELD, JR.	SENIOR COUNSEL	DEBORAH L. NAGLE, Ph.D.
SIBLEY P. REPPERT	JAMES E. COCKFIELD	ANNE JACQUELINE WIZEMAN, Ph.D.
DAVID R. BURNS	OF COUNSEL	BRIAN C. TRINQUE, Ph.D.
JOHN S. CURRAN	JEREMIAH LYNCH	CHRISTOPHER R. COWLES, Ph.D.
SEAN D. DETWEILER	JEANNE M. DIGIORGIO	JAMES H. VELEMA
MEGAN E. WILLIAMS, Ph.D.	CYNTHIA L. KANIK, Ph.D.	W. ELANA WANG
LISA DIROCCO TYNER	JOHN D. LANZA	CYNTHIA M. GILBERT
HATHAWAY P. RUSSELL *		
MARIA LACCOTRIPE ZACHARAKIS, Ph.D.		* Admitted in TX only
		** Admitted in CT only

September 29, 2004

MS Amendment
 Commissioner for Patents
 P.O. Box 1450
 Alexandria, Virginia 22313-1450

Re: U.S. Patent Application No.: 10/718,846
 For: *USE OF CREATINE OR CREATINE ANALOGS FOR THE TREATMENT OF DISEASES OF THE NERVOUS SYSTEM*
 Inventors: Kaddurah-Daouk, Rima *et al.*
 Filed: November 21, 2003
Our Ref. No.: AVZ-001CPUSCN

Dear Sir:

I enclose herewith for filing in the above-identified application the following:

1. Information Disclosure Statement;
2. PTO Form SB/08; and
3. A Return Postcard.

No additional costs are believed to be due in connection with the filing of this Information Disclosure Statement. However, please charge any necessary fees in connection with the enclosed statement to our Deposit Order Account No. 12-0080. For this purpose, a duplicate of this sheet is attached.

I hereby certify that this correspondence is deposited with the United States Postal Service as first class mail in an envelope addressed to:
 MS Amendment, Commissioner for Patents, P.O. Box 1450,
 Alexandria, Virginia 22313-1450 on:

September 29, 2004

Date



Meredith C. Arnold, Registration No. 52,568

Respectfully submitted,
 LAHIVE & COCKFIELD, LLP



Merideth C. Arnold
 Registration No. 52,568
 Attorney for Applicants



IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re the application of: Kaddurah-Daouk, Rima, *et al.*

Serial No.: 10/718,846

Filed: November 21, 2003

For: *USE OF CREATINE OR CREATINE ANALOGS FOR THE TREATMENT OF DISEASES OF THE NERVOUS SYSTEM*

Attorney Docket No.: AVZ-001CPUSCN

Group Art Unit: 1639

Examiner: Not Yet Assigned

MS Amendment
Commissioner for Patents
P.O. Box 1450
Alexandria, Virginia 22313-1450

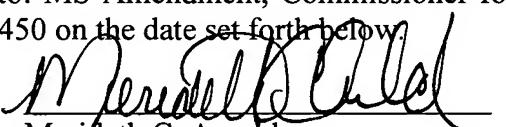
Certificate of First Class Mailing (37 CFR §1.8(a))

I hereby certify that this correspondence is being deposited with the United States Postal Service as first class mail in an envelope addressed to: MS Amendment, Commissioner for Patents, P.O. Box 1450, Alexandria, Virginia 22313-1450 on the date set forth below.

September 29, 2004

Date of Signature and of Mail Deposit

By:


Merideth C. Arnold
Registration No. 52,568
Attorney for Applicants

INFORMATION DISCLOSURE STATEMENT

Dear Sir:

For the Examiner's convenience in reviewing this continuation application, Applicants submit a consolidated PTO Form SB/08, listing all references cited during the prosecution of the parent application. The present application is a Continuation-in-Part of U.S. Serial No. 08/853,174, filed May 7, 1997 (Atty. Docket No. ATV-001CPUS). All references listed on the enclosed PTO Form SB/08 have been previously cited by or submitted to the Office in the prior application, and, in accordance with 37 CFR §1.98(d), copies of the references are not enclosed but will be provided upon request.

This statement is not to be interpreted as a representation that the cited publications are material, that an exhaustive search has been conducted, or that no other relevant information exists. Nor shall the citation of any publication herein be construed *per se* as a representation that such publication is prior art. Moreover, Applicants understand that the Examiner will make an independent evaluation of the cited publications.

Under 37 CFR § 1.97(b)(3), no additional costs are believed to be due in connection with the filing of this disclosure. If, however, a first Office Action on the merits issues in this application bearing a mailing date prior to the date of this Information Disclosure Statement, please charge the appropriate fee as required under 37 CFR §1.17(p) to our Deposit Order Account No. 12-0080.

Respectfully submitted,
LAHIVE & COCKFIELD, LLP

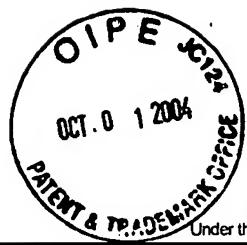


Merideth C. Arnold
Registration No. 52,568
Attorney for Applicants

28 State Street
Boston, MA 02109
(617) 227-7400

Date: September 29, 2004

EAH/CMS/MCA/r1
Enclosures



Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

PTO/SB/08a/b (08-03)

Approved for use through 07/31/2006. OMB 0651-0031

U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE

Substitute for form 1449A/B/PTO				Complete if Known	
				Application Number	10/718,846
				Filing Date	November 21, 2003
				First Named Inventor	Kaddurah-Daouk, Rima
				Art Unit	1639
				Examiner Name	Not Yet Assigned
Sheet	1	of	8	Attorney Docket Number	AVZ-001CPUSCN

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number Number-Kind Code ² (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
	A1	5,091,404	02-25-1992	Elgebaly, S.A.	
	A2	5,321,030	06-14-1994	Kaddurah-Daouk, R., et al.	
	A3	5,324,731	06-28-1994	Kaddurah-Daouk, R., et al.	
	A4	5,492,930	02-20-1996	Coffin, V.L.	
	A5	5,741,661	04-21-1998	Goldin, S.M., et al.	
	A6	6,169,115	01-02-2001	Kaddurah-Daouk, R., et al.	

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Foreign Patent Document Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
	A7	WO 90/09192	08-23-1990	Massachusetts Institute of Technology (US)	
	A8	WO 92/08456	05-29-1992	Amira Inc. (US)	
	A9	WO 94/16687	08-04-1994	University of Pennsylvania (US)	
	A10	WO 94/17794 A1	08-18-1994	Jennings, S.A. (GB)	

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. ¹Applicant's unique citation designation number (optional). ²See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶Applicant is to place a check mark here if English language Translation is attached.

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			
	A11	Annesley, T., et al., "Cyclocreatine Phosphate as a Substitute for Creatine Phosphate in Vertebrate Tissue. Energetic Considerations," <i>Biochem. Biophys. Res. Commun.</i> , 1977; 74:185-90.			
	A12	Annesley, T., et al., "Formation and Utilization of Novel High Energy Phosphate Reservoirs in Ehrlich Ascites Tumor Cells," <i>J. Biol. Chem.</i> , 1978; 253(22):8120-5.			
	A13	Beal, M., "Aging, Energy, and Oxidative Stress in Neurodegenerative Diseases," <i>Ann. Neurol.</i> , 1995; 38:357-66.			

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

Examiner Signature	Date Considered
--------------------	-----------------

Substitute for form 1449A/B/PTO				Complete if Known	
				Application Number	10/718,846
				Filing Date	November 21, 2003
				First Named Inventor	Kaddurah-Daouk, Rima
				Art Unit	1639
				Examiner Name	Not Yet Assigned
Sheet	2	of	8	Attorney Docket Number	AVZ-001CPUSCN

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number Number-Kind Code ² (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Foreign Patent Document Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
					T ⁶

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. ¹ Applicant's unique citation designation number (optional). ² See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			T ²
	B1	Beal, M., "Does Impairment of Energy Metabolism Result in Excitotoxic Neuronal Death in Neurodegenerative Illnesses?" <i>Ann. Neurol.</i> , 1992; 31:119-30.			
	B2	Beal, M. et al., "Age-Dependent Striatal Excitotoxic Lesions Produced by the Endogenous Mitochondrial Inhibitor Malonate," <i>J. Neurochem.</i> , 1993; 61:1147-50.			
	B3	Bertolini, et al., Chemical Abstracts AN 1982:49622.			
	B4	Brady, S., et al., "Nerve-Specific Enolase and Creatine Phosphokinase in Axonal Transport: Soluble Proteins and the Axoplasmic Matrix," <i>Cell</i> , 1981; 23:515-23.			
	B5	Brouillet, E., et al., "Chronic Mitochondrial Energy Impairment Produces Selective Striatal Degeneration and Abnormal Choreiform Movements in Primates," <i>PNAS</i> , 1995; 92:7105-9.			
	B6	Browne, S., et al., "Oxidative Damage and Metabolic Dysfunction in Huntington's Disease: Selective Vulnerability of the Basal Ganglia," <i>Ann. Neurol.</i> , 1997; 41:646-53.			
	B7	Burbaeva, G.S., et al., "Decreased Level of Immunoreactive Phosphokinase BB Isoenzymes in the Brain of Patients with Schizophrenia and Senile Dementia of the Alzheimer Type," <i>Zh. Nevropatol. Psichiatr Im S S Korsakova</i> , 1990; 90(7):85-7.			
	B8	Cadoux-Hudson, T., et al., "Imaging of Human Brain Creatine Kinase Activity in Vivo," <i>FASEB J.</i> , 1989; 3:2660-6.			

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

Examiner Signature	Date Considered
--------------------	-----------------

Substitute for form 1449A/B/PTO				<i>Complete if Known</i>	
				Application Number	10/718,846
				Filing Date	November 21, 2003
				First Named Inventor	Kaddurah-Daouk, Rima
				Art Unit	1639
				Examiner Name	Not Yet Assigned
Sheet	3	of	8	Attorney Docket Number	AVZ-001CPUSCN

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number Number-Kind Code ² (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Foreign Patent Document Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
					T ⁶

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. ¹ Applicant's unique citation designation number (optional). ² See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			T ²
	C1	Carter, A.J., et al., "Preincubation with creatine enhances levels of creatine phosphate and prevents anoxic damage in rat hippocampal slices," <i>J Neurochem.</i> 1995; 64(6):2691-9.			
	C2	Chandler, W., et al., "Regional Creatine Kinase, Adenylate Kinase, and Lactate Dehydrogenase in Normal Canine Brain," <i>Stroke</i> , 1988; 19:251-5.			
	C3	Cramer, F., et al., "Die Synthese der Argininphosphorsäure und die Reaktion von Isoureidophosphonaten mit Aminen," <i>Chem. Ber.</i> , 1962; 95:1670-82.			
	C4	De Leon, M., et al., "Identification of Transcriptionally Regulated Genes After Sciatic Nerve Injury," <i>J. Neurosci. Res.</i> , 1991; 29:437-48.			
	C5	Erecinska, M., et al., "ATP and Brain Function," <i>J. Cerebr. Blood Flow and Metabolism</i> , 1989; 9:2-19.			
	C6	Friedhoff, A., et al., "Creatine Kinase Isoenzyme Associated with Synaptosomal Membrane and Synaptic Vesicles," <i>Life Sci.</i> , 1977; 20:867-74.			
	C7	Goodman & Gilman, The Pharmacological Basis of Therapeutics," 1980, 6 th Ed. (MacMillan Pub.); 5-10.			
	C8	Griffiths, G., et al., "Accumulation of Analog of Phosphocreatine in Muscle of Chicks Fed 1-Carboxymethyl-2-iminoimidazolidine (Cyclocreatine)," <i>J. Biol. Chem.</i> , 1976; 251(7):2049-54.			

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

Examiner Signature	Date Considered
--------------------	-----------------

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449A/B/PTO				<i>Complete if Known</i>	
				Application Number	10/718,846
				Filing Date	November 21, 2003
				First Named Inventor	Kaddurah-Daouk, Rima
				Art Unit	1639
				Examiner Name	Not Yet Assigned
Sheet	4	of	8	Attorney Docket Number	AVZ-001CPUSCN

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number-Kind Code ² (if known)			

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Foreign Patent Document	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)			
					⁶

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. ¹ Applicant's unique citation designation number (optional). ² See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			
	D1	Gu, M., et al., "Mitochondrial Defect in Huntington's Disease Caudate Nucleus," <i>Ann. Neurol.</i> , 1996; 39:385-9.			
	D2	Gurney, M., et al., "Motor Neuron Degeneration in Mice That Express a Human Cu,Zn Superoxide Dismutase Mutation," <i>Science</i> , 1994; 264:1772-5.			
	D3	Hemmer, W., et al., "Functional Aspects of Creatine Kinase in Brain," <i>Dev. Neuroscience</i> , 1993; 15:249-260.			
	D4	Hagenfeldt, L., et al., "Creatine Treatment in MELAS," <i>Muscle and Nerve</i> . 1994; 10:1236-7.			
	D5	Hemmer, W., et al., "Creatine Kinase Isoenzymes in Chicken Cerebellum: Specific Localization of Brain-type Creatine Kinase in Bergmann Glial Cells and Muscle-type Creatine Kinase in Purkinje Neurons," <i>Eur. J. Neuroscience</i> , 1994; 6:538-49.			
	D6	Henshaw, R., et al., "Malonate Produces Striatal Lesions by Indirect NMDA Receptor Activation," <i>Brain Research</i> , 1994; 647:161-6.			
	D7	Hertz, L., et al., "Energy Metabolism at the Cellular Level of the CNS," <i>Can. J. Physiol. Pharmacol.</i> , 1992; 70:S145-57.			
	D8	Ito, M., "The Cellular Basis of Cerebellar Plasticity," <i>Corr. Opin. Neurobiol.</i> , 1991; 1:616-20.			
	D9	Jenkins, B., et al., "Evidence for Impairment of Energy Metabolism In Vivo in Huntington's Disease Using Localized ¹ H NMR Spectroscopy," <i>Neurology</i> , 1993; 43:2689-95.			

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

Examiner Signature	Date Considered
--------------------	-----------------

Substitute for form 1449A/B/PTO				Complete if Known	
				Application Number	10/718,846
				Filing Date	November 21, 2003
				First Named Inventor	Kaddurah-Daouk, Rima
				Art Unit	1639
				Examiner Name	Not Yet Assigned
Sheet	5	of	8	Attorney Docket Number	AVZ-001CPUSCN

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number Number-Kind Code ² (<i>if known</i>)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Foreign Patent Document Country Code ³ -Number ⁴ -Kind Code ⁵ (<i>if known</i>)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
					T ⁶

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. ¹ Applicant's unique citation designation number (optional). ² See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			
	E1	Kahn, M.A., "Effect of Calcium on Creatine Kinase Activity of Cerebellum," <i>Histochem.</i> , 1976; 48:29-32.			T ²
	E2	Lillie, J., et al., "Cyclocreatine (1-Carboxymethyl-2-iminoimidazolidine) Inhibits Growth of a Broad Spectrum of Cancer Cells Derived from Solid Tumors," <i>Cancer Res.</i> , 1993; 53:3172-8.			
	E3	Lim, L., et al., "Neurone-Specific Enolase and Creatine Phosphokinase are Protein Components of Rat Brain Synaptic Plasma Membranes," <i>J. Neurochem.</i> , 1983; 41:1177-82.			
	E4	LoPresti, P., et al., "Direct Determination of Creatine Kinase Equilibrium Constants with Creatine or Cyclocreatine as Substrate," <i>Biochem. Biophys. Acta.</i> , 1989; 998:317-20.			
	E5	Lowe, G., et al., "Evidence for an Associative Mechanism in the Phosphoryl Transfer Step Catalyzed by Rabbit Muscle Creatine Kinase," <i>J. Biol. Chem.</i> , 1980; 225(9):3944-51.			
	E6	Maker, H., et al., "Regional Changes in Cerebellar Creatine Phosphate Metabolism During Late Maturation," <i>Exp. Neurol.</i> , 1973; 38:295-300.			
	E7	Manos, P., et al., "Creatine Kinase Activity in Postnatal Rat Brain Development and in Cultured Neurons, Astrocytes, and Oligodendrocytes," <i>J. Neurochem.</i> , 1991; 56:2101-7.			
	E8	Marletta, M., et al., "Affinity Labeling of Creatine Kinase by N-(2,3-Epoxypropyl)-N-amidinoglycine," <i>J. Biol. Chem.</i> , 1979; 254(6):1879-86.			
	E9	McLaughlin, A., et al., "Specificity of Creatine Kinase for Guanidino Substrates," <i>J. Biol. Chem.</i> , 1972; 247(13):4382-8.			

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

Examiner Signature	Date Considered
--------------------	-----------------

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449A/B/PTO				Complete if Known	
				Application Number	10/718,846
				Filing Date	November 21, 2003
				First Named Inventor	Kaddurah-Daouk, Rima
				Art Unit	1639
				Examiner Name	Not Yet Assigned
Sheet	6	of	8	Attorney Docket Number	AVZ-001CPUSCN

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number Number-Kind Code ² (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Foreign Patent Document Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
					T ⁶

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. ¹ Applicant's unique citation designation number (optional). ² See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			T ²
	F1	Molloy, G., et al., "Rat Brain Creatine Kinase Messenger RNA Levels are High in Primary Cultures of Brain Astrocytes and Oligodendrocytes and Low in Neurons," <i>J. Neurochem.</i> , 1992; 59:1925-32.			
	F2	National Library of Medicine: IGM Metathesaurus Information Screen - Nervous System Diseases definitions and related terms.			
	F3	Newman, E., "Regulation of Potassium Levels by Glial Cells in the Retina," <i>Trends Neuroscience</i> , 1985; 8:156-9.			
	F4	Oblinger, M., et al., "Cytotypic Differences in the Protein Composition of the Axonally Transported Cytoskeleton in Mammalian Neurons," <i>J. Neurol.</i> , 1987; 7(2):453-62.			
	F5	Orlovskaia, D.D., et al., "Neuromorphology and Neurochemistry of Senile Dementias in the Light of Studies on Glial Response," <i>Vestn Ross Akad Med Nauk.</i> , 1992; 8:34-9.			
	F6	Rees, et al., Chemical Abstracts AN 89:82648, US patent 4,981,774.			
	F7	Reichenbach, A., "Glial K+ Permeability and CNS K+ Clearance by Diffusion and Spatial Buffering," <i>Acad. Sci. New York</i> , 1991; 272-86.			
	F8	Roberts, J., et al., "Higher Homolog and N-Ethyl Analog of Creatine as Synthetic Phosphagen Precursors in Brain, Heart, and Muscle, Repressors of Liver Amidinotransferase, and Substrates for Creatine Catabolic Enzymes," <i>J. Biol. Chem.</i> , 1985; 260(25):13502-8.			

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

Examiner Signature	Date Considered
--------------------	-----------------

Substitute for form 1449A/B/PTO				Complete if Known	
				Application Number	10/718,846
				Filing Date	November 21, 2003
				First Named Inventor	Kaddurah-Daouk, Rima
				Art Unit	1639
				Examiner Name	Not Yet Assigned
Sheet	7	of	8	Attorney Docket Number	AVZ-001CPUSCN

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number Number-Kind Code ² (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Foreign Patent Document Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
					T ⁶

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. ¹ Applicant's unique citation designation number (optional). ² See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³ Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴ For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵ Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶ Applicant is to place a check mark here if English language Translation is attached.

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			T ²
	G1	Roberts, J., et al., "Synthesis and Accumulation of an Extremely Stable High-Energy Phosphate Compound by Muscle, Heart, and Brain of Animals Fed the Creatine Analog, 1-Carboxyethyl-2-iminoimidazolidine (Homocyclocreatine)," <i>Arch. Biochem. Biophys.</i> , 1983; 220(2):563-71.			
	G2	Rowley, G.L., et al., "On the Specificity of Creatine Kinase. New Glycocyamines and Glycocyamine Analogs Related to Creatine," <i>Journal of the American Chemical Society</i> , 1971; 93:5542-51.			
	G3	Rudinger, "Peptide Hormones," 1976 (J. Parsons Ed.); 1-6.			
	G4	Schiffmann, R., et al., "Childhood Ataxia with Diffuse Central Nervous System Hypomyelination," <i>Ann. Neurol.</i> , 1994; 35:331-40.			
	G5	Schluthesis, et al., "Effects of Creatine on Synthesis and Release of Gamma-[3H] Aminobutyric Acid," <i>Journal of Neurochemistry</i> , 54(6):1858-63.			
	G6	Schultz, J., et al., "Blockade of Neuronal Nitric Oxide Synthase Protects Against Excitotoxicity in vivo," <i>J. Neurosci.</i> , 1995; 15(12):8419-29.			
	G7	Schultz, J., et al., "Inhibition of Neuronal Nitric Oxide Synthase by 7-Nitroindazole Protects Against MPTP-Induced Neurotoxicity in Mice," <i>J. Neurochem.</i> , 1995; 64:936-9.			

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

Examiner Signature	Date Considered
--------------------	-----------------

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it contains a valid OMB control number.

Substitute for form 1449A/B/PTO				<i>Complete if Known</i>	
				Application Number	10/718,846
				Filing Date	November 21, 2003
				First Named Inventor	Kaddurah-Daouk, Rima
				Art Unit	1639
				Examiner Name	Not Yet Assigned
Sheet	8	of	8	Attorney Docket Number	AVZ-001CPUSCN

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Document Number	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number-Kind Code ² (if known)			

FOREIGN PATENT DOCUMENTS					
Examiner Initials*	Cite No. ¹	Foreign Patent Document	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Country Code ³ -Number ⁴ -Kind Code ⁵ (if known)			
					T ⁶

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. ¹Applicant's unique citation designation number (optional). ²See Kinds Codes of USPTO Patent Documents at www.uspto.gov or MPEP 901.04. ³Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). ⁴For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. ⁵Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST.16 if possible. ⁶Applicant is to place a check mark here if English language Translation is attached.

NON PATENT LITERATURE DOCUMENTS					
Examiner Initials*	Cite No. ¹	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.			
	H1	Stadhouders, A., et al., "Mitochondrial Creatine Kinase: A Major Constituent of Pathological Inclusions Seen in Mitochondrial Myopathies," PNAS, 1994; 91(11):5089-93.			
	H2	Wang, T., "Synthesis and Properties of N-Acetimidoyl Derivatives of Glycine and Sarcosine," JOC, 1974; 39(24):3591-4.			

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

¹Applicant's unique citation designation number (optional). ²Applicant is to place a check mark here if English language Translation is attached.

Examiner Signature	Date Considered
--------------------	-----------------